Appl. No. 10/559,878 Amdt. Dated December 3, 2010

Reply to Office Action of September 3, 2010

## Amendments to the Claims

This listing of claims will replace all prior versions, and listings, of claims in the application:

## Listing of Claims:

 (Currently amended) A method for the production of an aromatic or hetroaromatic fluorine-labelled compound comprising fluoridation of an iodonium salt of Formula (I) or (ID:

wherein:

Q = phenyl, or phenyl substituted with one or more of acetyl, C1-3 alkyl or -0-C1-3 alkyl each of  $R^4$ ,  $R^3$ ,  $R^4$  and  $R^5$  is independently hydrogen,  $O(C_{L+0}$  alkyl) or  $C_{L+0}$  alkyl or protected versions thereof  $R^1$ - $R^5$  is independently selected from hydrogen,  $C_{L-3}$  alkyl and -0- $(C_1$ - $C_3$  alkyl); and

 $Y^{-}$  is a counter ion such as trifluoromethane sulfonate (triflate), perfluoro  $C_2$ - $C_{10}$  alkyl sulphonate, trifluoroacetate, methane sulfonate (mesylate), toluene sulfonate. (tosylate), tetraphenylborate;

to give a product of general formula (III):

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where Q is substituted with one or more substituents selected from  $C_{++0}$  alkyl,  $O(C_{++0}$  alkyl), C(=0)  $C_{++0}$  alkyl), C(=0) C(

 $O(C_{5,14} \text{ aryl})$ ,  $C(=O)C_{5,14} \text{ aryl}$ ,  $C(=O)NR^6(C_{5,14} \text{ aryl}$ ,  $C_{5,14} \text{ heteroaryl}$ ,  $O(C_{5,14} \text{ heteroaryl})$ .

C(=0)C<sub>5-14</sub> heteroaryl, C(=0)NR<sup>6</sup>(C<sub>5-14</sub> heteroaryl), C<sub>3-16</sub> cycloalkyl, O(C<sub>3-16</sub> cycloalkyl),

 $C(=O)(C_{3-10}-cycloalkyl)$ ,  $C(=O)NR^{6}(C_{3-10}-cycloalkyl)$ ,  $C_{3-10}-heterocyclyl$ ,  $O(C_{3-10}-cycloalkyl)$ 

heterocyclyl),  $C(=0)(C_{3.40}$  heterocyclyl),  $C(=0)NR^6(C_{5.44}$  heterocyclyl) or protected

versions thereof, when Q is substituted with an electron donating substituent, Q also contains

one or more electron withdrawing groups to ensure Q is electron deficient;

and wherein said fluoridation is carried out with a fluoride ion source characterised in that

the reaction solvent is either 100% water or a mixture of water and a water miscible solvent.

(Cancelled)

(Cancelled)

4. (Previously presented) A method as claimed in claim 1, wherein the water miscible

 $solvent\ is\ acetonitrile,\ ethanol,\ methanol,\ tetrahydrofuran\ or\ dimethyl formamide.$ 

5. (Previously Presented) A method as claimed in claim 1 wherein the volume:volume

ratio of water:water-miscible solvent is between 1:99 and 1:1.

6. (Original) A method as claimed in claim 5 wherein the volume:volume ratio of

water:water-miscible solvent is from 10:90 to 30:70.

7. (Previously Presented) A method as claimed in claim 1, wherein the fluoride ion

source is potassium, caesium or sodium fluoride.

8. (Cancelled)

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(Cancelled)

 $10. \hspace{0.5cm} \hbox{(Previously Presented)} \hspace{0.5cm} \hbox{A method as claimed in claim 1 wherein, in the compound} \\$ 

of Formula II, the "solid support" is polystyrene, polyacrylamide, polypropylene or glass

or silicon coated with such a polymer.

11. (Previously Presented) A method as claimed in claim 1 wherein the solid support is in

the form of small discrete particles or is a coating on the inner surface of a reaction vessel.

12. (Previously Presented) A method as claimed in claim 1, wherein, in the compound of

Formula II the "linker" is  $C_{1\text{--}20}$  alkyl or  $C_{1\text{--}20}$  alkoxy, attached to the resin by an amide ether

or a sulphonamide bond or a polyethylene glycol (PEG) linker.

13. (Previously Presented) A method as claimed in claim 1

wherein  $R^6$  is H,  $C_1\text{-}C_6$  alkyl,  $C_3\text{-}C_{10}$  cycloalkyl,  $C_3\text{-}C_{10}$  heterocyclyl,  $C_4\text{-}C_{10}$  aryl or

C<sub>4</sub>-C<sub>10</sub> heteroaryl;

any of which may optionally be substituted with OH, NHR<sup>6</sup>, COOH or protected versions

any of these groups; or alternatively

any two adjacent substituents may form a four- to six-membered carbocyclic or heterocyclic

ring, optionally fused to a further aromatic, heteroaromatic, carbocyclic or heterocyclic ring.

14-15. (Cancelled)

16. (Previously Presented) A method as claimed in claim 1, wherein the fluorine-labelled

compound is an [18F]-labelled compound and the fluoride ion source is a source of 18F.

17. (Cancelled)

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18. (Previously Presented) A method as claimed in claim 1, further including, in any order, one or more of the following steps: removal of excess <sup>18</sup>F̄, for example by ion-exchange chromatography; and/or

- (i) removal of the protecting groups; and/or
- (ii) removal of organic solvent; and/or
- (iii) formulation of the resultant compound as an aqueous solution.

19. (Previously Presented) A kit for the production of an aromatic fluorine-labelled compound, the kit comprising:

- (i) a vial containing an aqueous solvent for dissolving the fluoride ion source; and
- (ii) a reaction vessel containing an iodonium salt of claim 1.
- 20. (Original) A kit as claimed in claim 19, wherein the solvent is 100% water.

 (Original) A kit as claimed in claim 19 wherein the solvent is a mixture of water and a water miscible solvent.

- 22. (Original) A kit as claimed in claim 21, wherein the water miscible solvent is acetonitrile, ethanol, methanol, tetrahydrofuran or dimethylformamide.
- 23. (Previously Presented) A kit as claimed in claim 21 wherein the volume:volume ratio of water:water-miscible solvent is between 1:99 and 1:1.
- 24. (Original) A kit as claimed in claim 23 wherein the volume:volume ratio of water-water-miscible solvent is from 10:90 to 30:70.
- 25-26. (Cancelled)

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- 27. (Previously Presented) A kit as claimed in claim 19, wherein the reaction vessel is a cartridge or a microfabricated vessel.
- 28. (Previously Presented) A kit as claimed in claim 19, further comprising a source of fluoride ions.